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21) International Application Number: PCT/US 22) International Filing Date: 1 November 1999 (30) Priority Data: 60/106,732 2 November 1998 (02.11.98 31) Applicant (for all designated States except US): TF ERNMENT OF THE UNITED STATES OF AN represented by THE SECRETARY OF THE I MENT OF HEALTH AND HUMAN SERVICES Bethesda, MD 20892 (US). 32) Inventors; and 35) Inventors; and (US). NEWTON, Dianne, L. [US/US]; 15904 New Drive, Rockville, MD 20855 (US). 34) Agents: WEBER, Kenneth, A. et al.; Townsend and T and Crew LLP, 8th floor, Two Embarcadero Ce Francisco, CA 94111–3834 (US).	O1.11.99 DE GOV MERICA DEPART [US/US anna, M D 2170 Bedfor	BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JF KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM GA, GN, GW, ML, MR, NE, SN, TD, TG). Published With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: SELECTIVE TOXICITY OF AMINO-TERMINAL MODIFIED RNASE A SUPERFAMILY POLYPEPTIDES

(57) Abstract

This invention provides RNase A superfamily polypeptides with modified amino terminal which can be used to selectively kill target Kaposi's sarcoma cells, neoplastic endothelial cells, and non-neoplastic endothelial cells. In certain embodiments of the invention, the amino terminal modification consists of an addition of 4 amino acid sequence consisting of the SLHV sequence at position -4 to -1 to the eosinophil derived neurotoxin protein. The amino terminal addition is capable of directing the claimed RNase A superfamily polypeptides to proliferating endothelial cells, such as Kaposi's sarcoma cells, and selectively killing these cells.

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